

RAW SEQUENCE LISTING DATE: 07/09/2002 PATENT APPLICATION: US/09/705,985 TIME: 16:07:59

Input Set : A:\2874-B.ST25.txt

Output Set: N:\CRF3\07092002\1705985.raw

PECEIVED

3 <110> APPLICANT: ANDERSON, Dirk, M. JUL 1 9 2002 4 GALIBERT, Laurent, J. 6 <120> TITLE OF INVENTION: METHOD OF INHIBITING OSTEOCLAST ACTIVITY 8 <130> FILE REFERENCE: 2874-B JIT OCHTER 1600/2900 10 <140> CURRENT APPLICATION NUMBER: 09/705,985 11 <141> CURRENT FILING DATE: 2000-11-03 13 <150> PRIOR APPLICATION NUMBER: PCT/US99/10588 14 <151> PRIOR FILING DATE: 1999-05-13 ENTERED 16 <150> PRIOR APPLICATION NUMBER: 60/085,487 17 <151> PRIOR FILING DATE: 1998-05-14 19 <150> PRIOR APPLICATION NUMBER: 60/110,836 20 <151> PRIOR FILING DATE: 1998-12-03 22 <150> PRIOR APPLICATION NUMBER: 08/996,139 23 <151> PRIOR FILING DATE: 1997-12-22 25 <150> PRIOR APPLICATION NUMBER: 60/064,671 26 <151> PRIOR FILING DATE: 1997-10-14 28 <150> PRIOR APPLICATION NUMBER: 60/077,181 29 <151> PRIOR FILING DATE: 1997-03-07 31 <150> PRIOR APPLICATION NUMBER: 60/059,978 32 <151> PRIOR FILING DATE: 1996-12-23 34 <160> NUMBER OF SEQ ID NOS: 8 36 <170> SOFTWARE: PatentIn version 3.1 38 <210> SEQ ID NO: 1 39 <211> LENGTH: 3136 40 <212> TYPE: DNA 41 <213> ORGANISM: Homo sapiens 43 <220> FEATURE: 44 <221> NAME/KEY: CDS 45 <222> LOCATION: (39)..(1886) 46 <223> OTHER INFORMATION: 49 <400> SEQUENCE: 1 50 cegetgagge egeggegee geeageetgt eeegegee atg gee eeg ege ege 56 51 Met Ala Pro Arg Ala Arg 52 104 55 Arg Arg Arg Pro Leu Phe Ala Leu Leu Leu Cys Ala Leu Leu Ala 10 58 cgg ctg cag gtg gct ttg cag atc gct cct cca tgt acc agt gag aag 152 59 Arg Leu Gln Val Ala Leu Gln Ile Ala Pro Pro Cys Thr Ser Glu Lys 30 200 62 cat tat gag cat ctg gga cgg tgc tgt aac aaa tgt gaa cca gga aag 63 His Tyr Glu His Leu Gly Arg Cys Cys Asn Lys Cys Glu Pro Gly Lys

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66 tac 67 Tyr 68 55	_			Lys	_					-	-	_		_		248
70 tgt 71 Cys 72				gaa	tac			Ser		aat					tgc	296
74 ttg 75 Leu 76				gtt				ggc	aag					gtg		344
78 gcc 79 Ala 80																392
82 cac 83 His 84																440
86 ccg 87 Pro 88 135		_		Ala	_		-	-				-				488
90 tgc 91 Cys 92			Cys		-			Phe		-	-				_	536
94 gac 95 Asp 96		-	_					_					_	_	-	584
98 gaa	cat.	cat	ααα	202	παπ	222	too	αat	aca	att	tac	~~+	+ -+	+ a+	a+ a	632
99 Glu 100			Gly					Asp		-	_	_	Ser		-	032
99 Glu	His gct	His 185 aga Arg	Gly Gly Gly Gly	Thr cca	Glu cca	Lys aat	Ser 190 gaa Glu	Asp	Ala cat	Val gtt	Cys tac	Ser 195 ttg Leu	Ser ; , cc	Ser ggt	Leu tta	680
99 Glu 100 102 cca 103 Pro	His gct Ala 200 att	His 185 aga Arg	Gly Gly a aaa J Lys	Thr cca Pro ctc	Glu cca Pro	Lys aat Asn 205 gcg	Ser 190 gaa Glu	Asp ccc Pro	Ala cat His	Val gtt Val	Cys tac Tyr 210 gtg	Ser 195 ttg Leu) g gct	Ser ; ; ccc ; Pro	Ser ggt Gly	Leu tta Leu atc	
99 Glu 100 102 cca 103 Pro 104 106 ata 107 Ile	His gct Ala 200 a att	His 185 1 aga 1 Arg 1 ctg 2 ctg 2 teg 2 gtt	Gly i aaa j Lys ctt Leu tgc	Thr cca Pro ctc Leu tat	CCa Pro tto Phe 220 agg	Lys aat Asn 205 gcg Ala	Ser 190 gaa Glu tct Ser	Asp ccc Pro gtg Val	cat His gcc Ala	Val gtt Val ctg Leu 225	Cys tac Tyr 210 ggtg Val	Ser 195 ttg Leu) gct Ala	Ser	Ser ggt Gly ato Ile	Leu tta Leu atc Ile 230 ttg Leu	680
99 Glu 100 102 cca 103 Pro 104 106 ata 107 Ile 108 215 110 ttt 111 Phe 112 114 tgg 115 Trg 116	His gct 200 Ala 200 atte Ile Gly Caco His	His 185 aga Arg Ctg Leu gtt Val	Gly aaaa J Lys ctt Leu Cys Cys atc J Ile 250	Thr cca Pro ctc Leu tat Tyr 235 aat Asn	cca Pro ttc Phe 220 agg Arg gag Glu	Lys aat Asn 205 gcg Ala aaa Lys gct Ala	Ser 190 gaa Glu tct Ser aaa Lys	Asp ccc Pro gtg Val ggg Gly ggc Gly 255	cat His gcc Ala aaa Lys 240 cgc	Val gtt Val ctg Leu 225 gca Ala	Cys tac Tyi 210 ygto Val ctc Leu agt	Ser 195 c ttg c ttg c Leu) g gct l Ala c aca c Thr c gga	Ser g cco g pro a gct a gct a Ala gat gat gat gat gat	Ser ggt ggt atca lle aat aAsn 245 aag Lys	tta Leu atc Ile 230 ttg Leu gag Glu	680 728
99 Glu 100 102 cca 103 Pro 104 106 ata 107 Ile 108 215 110 ttt 111 Phe 112 114 tgg 115 Trg	His A gct Ala 200 A att Graph Graph Graph Graph Gra	His 185 aga Arg Core Core Core Core Core Core Core Core	Gly aaaa Lys ctt Leu Cys atc 250 gac Asp	Thr cca Pro ctc Leu tat Tyr 235 aat Asn	cca Pro ttc Phe 220 agg Arg gag Glu	Lys aat Asn 205 gcg Ala aaa Lys gct Ala	Ser 190 gaa Glu tct Ser aaa Lys tgt	Asp ccc Pro gtg Val ggg Gly 255 aca Thr	cat His gcc Ala aaa Lys 240 cgc Arg	Val gtt Val ctg Leu 225 gca Ala cta Leu	Cys tac Tyr 210 ygto Val cto Leu ser gca	Ser 195 2 ttg 2 Leu) g gct Ala 2 aca 1 Thr 2 gga 2 Gly	Ser G CCC G Pro Ala G Ala G Ala Asr Asr 260 C ttt Phe	Ser ggt ggt atcalle attal Asn 245 aag Lys ggt	Leu tta Leu atc Ile 230 ttg Leu gag Glu cag	680 728 776
99 Glu 100 102 cca 103 Pro 104 106 ata 107 Ile 108 215 110 ttt 111 Phe 112 114 tgg 115 Trg 116 118 tcc 119 Ser	His get Ala 200 att Get Get Get Get Get Get Get	His 185 aga Argo Cto Cto Cto Cto Cto Cto Cto Cto Cto Ct	Gly aaaa Lys ctt Leu Cys atc 250 gac Asp tgt	Thr cca Pro ctc Leu tat Tyr 235 aat Asn agt Ser	cca Pro ttc Phe 220 agg Arg Glu tgt Cys	Lys aat Asn 205 gcg Ala aaa Lys gct Ala gtc	Ser 190 gaa Glu tct Ser aaa Lys tgt Cys agt 270 tta Leu	Asp ccc Pro gtg Val ggg Gly 255 aca Thr	cat His gcc Ala aaa Lys 240 cgc Arg cac	Val gtt Val ctg Leu 225 gca Ala cta Leu acg	Cys tac Tyr 210 yeta tac tac tac tac tac tac tac	Ser 195 c ttg c ttg c tag d Ala c aca d Thr c gga d Asn 275 gag d Glu	Ser Grant Pro Grant Ala Grant Ala Grant Asp 260 Crant Phe	Ser ggt catcalle aata Asn 245 aag Lys ggt eGly aag	Leu tta Leu atc Ile 230 ttg Leu gag Glu cag Gln aca	680 728 776 824
99 Glu 100 102 cca 103 Pro 104 106 ata 107 Ile 108 215 110 ttt 111 Phe 112 114 tgg 115 Trg 116 118 tcc 119 Ser 120 122 cag 123 Glr	His A gct Ala 200 Ata 200 Ata Constant Grade Gly Gly Grade Gly Gly Gly Constant Grade Gly Constant Gly	His 185 aga a Argo cto cto cto cto cto cto cto cto cto ct	Gly aaaa Lys ctt Leu tgc Cys atc Jac gac Asp tgt Cys	Thr cca Pro ctc Leu tat Tyr 235 aat Asn agt Ser gaa Glu atg	cca Pro ttc Phe 220 agg Arg Glu tgt Cys	Lys aat Asn 205 gcg Ala aaa Lys gct Ala gtc Val 285 tac Tyr	ser 190 gaa Glu tct ser aaa Lys tgt Cys agt 270 tta Leu	Asp ccc pro gtg Val ggg Gly 255 aca Thr ctg Leu	cat His gcc Ala aaa Lys 240 cgc Arg cac His ctg	Val gtt Val ctg Leu 225 gca Ala cta Leu acg Thr	Cys tac Tyr 210 gtg Val ctc Leu Ala ctg ggt Gly	Ser 195 c ttg 19	Ser Grant Pro Grant Ala Grant Gat Grant	Ser ggt catcalle aatta Asn 245 caag b Lys cag t Lys cag cag cag cag	Leu tta Leu atc Ile 230 ttg Leu gag Glu cag Gln aca Thr	680 728 776 824 872

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		_					_	_					_		_		
	Thr	Cys	Val	GLy	_	Gly	Pro	Tyr	Ala		Gly	Glu	Asp	Ala		Met	
132					315					320					325		1004
	ctc																1064
	Leu	Ser	Leu		Ser	rys	Thr	GIU		GIU	GLU	Asp	ser		Arg	GIN	
136				330					335					340			1110
	atg			-	_	-		-	-	-			_			-	1112
	Met	Pro		GIu	Asp	Glu	Tyr		Asp	Arg	Pro	Ser		Pro	Thr	Asp	
140			345					350					355				1160
	cag																1160
	Gln		Leu	Phe	Leu	Thr		Pro	GLY	Ser	Lys		Thr	Pro	Pro	Phe	
144		360					365					370					
	tct																1208
	Ser	Glu	Pro	Leu	Glu		Gly	Glu	Asn	Asp		Leu	Ser	Gln	Cys		
	375					380					385					390	
	acg																1256
	Thr	Gly	Thr	Gln		Thr	Val	Gly	Ser		Ser	Cys	Asn	Cys		Glu	
152					395					400					405		
	ccc																1304
	Pro	Leu	Cys	-	Thr	Asp	\mathtt{Trp}	Thr		Met	Ser	Ser	Glu		Tyr	Leu	
156				410					415					420			
	caa																1352
159	Gln	Lys	Glu	Val	Asp	Ser	Gly		Cys	Pro	His	Trp	Ala	Ala	Ser	Pro	
160			425					430					435				
	agc				-	-	-	_			_						1400
163	Ser	Pro	Asn	\mathtt{Trp}	Ala	Asp	Val	Cys	Thr	Gly	Cys	Arg	Asn	Pro	Pro	Gly	
164		440					445					450					
	gag																1448
167	Glu	Asp	Cys	Glu	Pro	Leu	Val	Gly	Ser	Pro	Lys	Arg	Gly	Pro	Leu	Pro	
	455					460					465					470	
	cag																1496
	Gln	Cys	Ala	Tyr	Gly	Met	Gly	Leu	Pro	Pro	Glu	Glu	Glu	Ala	Ser	Arg	
172					475					480					485		
174	acg	gag	gcc	aga	gac	cag	ccc	gag	gat	ggg	gct	gat	ggg	agg	ctc	cca	1544
175	Thr	Glu	Ala	Arg	Asp	Gln	Pro	Glu	Asp	Gly	Ala	Asp	Gly	Arg	Leu	Pro	
176				490					495					500			
178	agc	tca	gcg	agg	gca	ggt	gcc	ggg	tct	gga	agc	tcc	cct	ggt	ggc	cag	1592
179	Ser	Ser	Ala	Arg	Ala	Gly	Ala	Gly	Ser	Gly	Ser	Ser	${\tt Pro}$	Gly	Gly	Gln	
180			505					510					515				
182	tcc	cct	gca	tct	gga	aat	gtg	act	gga	aac	agt	aac	tcc	acg	ttc	atc	1640
183	Ser	${\tt Pro}$	Ala	Ser	Gly	Asn	Val	Thr	Gly	Asn	Ser	Asn	Ser	Thr	Phe	Ile	
184		520					525					530					
186	tcc	agc	ggg	cag	gtg	atg	aac	ttc	aag	ggc	gac	atc	atc	gtg	gtc	tac	1688
187	Ser	Ser	Gly	Gln	Val	Met	Asn	Phe	Lys	Gly	Asp	Ile	Ile	Val	Val	Tyr	
188						540					545					550	•
	gtc																1736
	Val																
192					555					560					565		
194	ggc	cgc	ccg	gtg	cag	gag	gag	acc	ctg	gcg	cgc	cga	gac	tcc	ttc	gcg	1784
	Gly																
											_						

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196 570 575 580	
198 ggg aac ggc ccg cgc ttc ccg gac ccg tgc ggc ggc ccc gag ggg ctg	1832
199 Gly Asn Gly Pro Arg Phe Pro Asp Pro Cys Gly Gly Pro Glu Gly Leu	
200 585 590 595	
202 cgg gag ccg gag aag gcc tcg agg ccg gtg cag gag caa ggc ggg gcc	1880
203 Arg Glu Pro Glu Lys Ala Ser Arg Pro Val Gln Glu Gln Gly Ala	
204 600 605 610	
206 aag get tgagegeece ceatggetgg gageeegaag eteggageea gggetegega	1936
207 Lys Ala	
208 615	
210 gggcagcacc gcagcctctg ccccagcccc ggccacccag ggatcgatcg gtacagtcg	1996
212 ggaagaccac ccggcattct ctgcccactt tgccttccag gaaatgggct tttcaggaag	g 2056
214 tgaattgatg aggactgtcc ccatgcccac ggatgctcag cagcccgccg cactggggca	a 2116
216 gatgtctccc ctgccactcc tcaaactcgc agcagtaatt tgtggcacta tgacagcta	2176
218 ttttatgact atcctgttct gtgggggggg ggtctatgtt ttccccccat atttgtattc	2236
220 cttttcataa cttttcttga tatctttcct ccctcttttt taatgtaaag gttttctcaa	a 2296
222 aaatteteet aaaggtgagg gtetetttet tttetettt cettttttt ttettttt	2356
224 ggcaacctgg ctctggccca ggctagagtg cagtggtgcg attatagccc ggtgcagcc	2416
226 ctaactcctg ggctcaagca atccaagtga tcctcccacc tcaaccttcg gagtagctgg	2476
228 gatcacaget geaggeeacg eccagettee tecceeegae tecceeece eagagaeac	g 2536
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234 ttcttttgtg cccctgctca cagtgtttta gagatggctt tcccagtgtg tgttcattg	
236 aaacactttt gggaaagggc taaacatgtg aggcctggag atagttgcta agttgctagg	g 2776
238 aacatgtggt gggactttca tattctgaaa aatgttctat attctcattt ttctaaaaga	
240 aagaaaaaag gaaacccgat ttatttctcc tgaatctttt taagtttgtg tcgttcctta	
242 agcagaacta agctcagtat gtgaccttac ccgctaggtg gttaatttat ccatgctgg	
244 agaggcactc aggtacttgg taagcaaatt tctaaaactc caagttgctg cagcttggca	
246 ttcttcttat tctagaggtc tctctggaaa agatggagaa aatgaacagg acatggggc	3076
248 cctggaaaga aagggcccgg gaagttcaag gaagaataaa gttgaaattt taaaaaaaaa	3136
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10 1 E 10 1E	
259 1 5 10 15	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263 20 25 30	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263 20 25 30 266 Pro Cys Thr Ser Glu Lys His Tyr Glu His Leu Gly Arg Cys Cys Asn	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263 20 25 30 266 Pro Cys Thr Ser Glu Lys His Tyr Glu His Leu Gly Arg Cys Cys Asn 267 35 40 45	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263 20 25 30 266 Pro Cys Thr Ser Glu Lys His Tyr Glu His Leu Gly Arg Cys Cys Asn 267 35 40 45 270 Lys Cys Glu Pro Gly Lys Tyr Met Ser Ser Lys Cys Thr Thr Thr Ser	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263 20 25 30 266 Pro Cys Thr Ser Glu Lys His Tyr Glu His Leu Gly Arg Cys Cys Asn 267 35 40 45 270 Lys Cys Glu Pro Gly Lys Tyr Met Ser Ser Lys Cys Thr Thr Thr Ser 271 50 55 60	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263 20 25 30 266 Pro Cys Thr Ser Glu Lys His Tyr Glu His Leu Gly Arg Cys Cys Asn 267 35 40 45 270 Lys Glu Pro Gly Lys Tyr Met Ser Ser Lys Cys Thr Thr Thr Ser 271 50 55 60 274 Asp Ser Val Cys Leu Pro Cys Gly Pro Asp Glu Tyr Leu Asp Ser Trp 275 65 70 75	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263	
262 Leu Cys Ala Leu Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 25 30	
262 Leu Cys Ala Leu Leu Ala Arg Leu Gln Val Ala Leu Gln Ile Ala Pro 263	

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286 287		Cys	Thr 115	Ala	Gly	Tyr	His	Trp 120	Ser	Gln	Asp	Cys	Glu 125	Cys	Cys	Arg
290 291	-	Asn 130	Thr	Glu	Cys	Ala	Pro 135	Gly	Leu	Gly	Ala	Gln 140	His	Pro	Leu	Gln
	Leu 145	Asn	Lys	Asp	Thr	Val 150	Cys	Lys	Pro	Cys	Leu 155	Ala	Gly	Tyr	Phe	Ser 160
298 299	Asp	Ala	Phe	Ser	Ser 165	Thr	Asp	Lys	Cys	Arg 170	Pro	Trp	Thr	Asn	Cys 175	Thr
	Phe	Leu	Gly	Lys 180	Arg	Val	Glu	His	His 185	Gly	Thr	Glu	Lys	Ser 190	Asp	Ala
306 307		Суѕ	Ser 195	Ser	Ser	Leu	Pro	Ala 200	Arg	Lys	Pro	Pro	Asn 205	Glu	Pro	His
	Val	Tyr 210	Leu	Pro	Gly	Leu	Ile 215		Leu	Leu	Leu	Phe 220		Ser	Val	Ala
314			Ala	Ala	Ile	Ile 230		Gly	Val	Cys	Tyr 235		Lys	Lys	Gly	Lys 240
		Leu	Thr	Ala	Asn 245		Trp	His	Trp	Ile 250		Glu	Ala	Cys	Gly 255	
	Leu	Ser	Gly	Asp 260		Glu	Ser	Ser	Gly 265		Ser	Cys	Val	Ser 270		His
	Thr	Ala	Asn 275		Gly	Gln	Gln	Gly 280		Cys	Glu	Gly	Val 285	Leu	Leu	Leu
	Thr	Leu 290		Glu	Lys	Thr	Phe 295		Glu	Asp	Met	Cys		Pro	Asp	Gln
334	Gly 305		Val	Cys	Gln	Gly 310		Cys	Val	Gly	Gly 315		Pro	Tyr	Ala	Gln 320
		Glu	Asp	Ala	Arg 325		Leu	Ser	Leu	Val 330		Lys	Thr	Glu	Ile 335	
	Glu	Asp	Ser	Phe 340		Gln	Met	Pro	Thr 345		Asp	Glu	Tyr	Met 350		Arg
	Pro	Ser	Gln 355		Thr	Asp	Gln	Leu 360		Phe	Leu	Thr	Glu 365	Pro	Gly	Ser
350 351	Lys	Ser 370	Thr	Pro	Pro	Phe	Ser 375	Glu	Pro	Leu	Glu	Val 380	Gly	Glu	Asn	Asp
	Ser 385	Leu	Ser	Gln	Cys	Phe 390	Thr	Gly	Thr	Gln	Ser 395	Thr	Val	Gly	Ser	Glu 400
358 359	Ser	Cys	Asn	Cys	Thr 405	Glu	Pro	Leu	Cys	Arg 410	Thr	Asp	Trp	Thr	Pro 415	Met
362 363					_			_			_		_	His 430	_	Pro
366 367	His	Trp	Ala 435	Ala	Ser	Pro	Ser	Pro 440	Asn	Trp	Ala	Asp	Val 445	Cys	Thr	Gly
370 371	Cys	Arg 450	Asn	Pro	Pro	Gly	Glu 455	Asp	Cys	Glu	Pro	Leu 460	Val	Gly	Ser	Pro
	Lys 465	Arg	Gly	Pro	Leu	Pro 470		Cys	Ala	Tyr	Gly 475		Gly	Leu	Pro	Pro 480
		Glu	Glu	Ala	Ser 485	Arg	Thr	Glu	Ala	Arg 490		Gln	Pro	Glu	Asp 495	Gly
	Ala	Asp	Gly	Arg		Pro	Ser	Ser	Ala		Ala	Gly	Ala	Gly		Gly

VERIFICATION SUMMARY

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